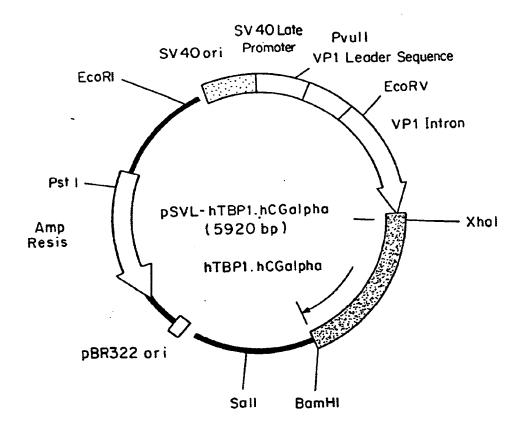


6663867

F1G. 1a(1)





F16.10(2)

PCU Interes

ATG GCT ACA G <u>gtaagcgccctaaaatccctttgggcacaatgtgtctgaggggagaggggagcgctgtagatgggacggggggactaagctttggggtttct</u> • Met Ala Th*t* hGH Signal Sequence

Pro Pro CTG CTC CTG 63C CrG GCT TTT Leu Ala Phe CTC TCC CTG Thr CGGCTCCCTCTGTTGCCCTCTGGTTTCTCCCCCAGGC TCC CGG +20 Asp of Processed TBP1

CAA GAG GGC AGT GCC GAT AGT GTG TGT CCC CAA GGA AAA TAT ATC CAC CCT CAA AAT AAT TCG ATT TGC TGT ACC AAG TGC CAC AAA AAA GIN GIU Gly Ser Ala Aap ser val Cys Pro Gln Gly Lys Tyr Ile His Pro Gln Asn Asn Ser Ile Cys Cys Thr Lys Cys His Lys Gly

ACC TAC TTG TAC AAT GAC TGT CCA GGC CCG GGG CAG GAT ACG GAC TGC AGG GAG TGT GAG AGC GGC TCC TTC ACC GCT TCA GAA AAC CAC CTC

GGTCAG GAG AAA CAG AAC ACC GTG TGC TGC CAT GCA GGT TTC TTT CTA AGA GAA AAC GAG TGT GTC TCC TGT GCC GGT GCT GCC CCA Thr Cys His Ala Gly Phe Phe Leu Arg Glu Asn Glu Cys Val Ser Cys Ala Gly Ala Ala Gln Glu Lys Gln Asn Thr Val Cys

TGC CCA GAA TGC ACG CTA CAG GAA AAC CCA TTC TTC TCC CAG CCG GGT GCC CCA ATA CTT CAG TGC ATG GGC TGC TTC TCT AGA GCA TAT Cys Pro Glu Cys Thr Leu Gln Glu Asn Pro Phe Phe Ser Gln Pro Gly Ala Pro Ile Leu Gln Cys Met Gly Cys Cys Phe Ser Arg Ala Tyr +7 Cys of hCG alpha

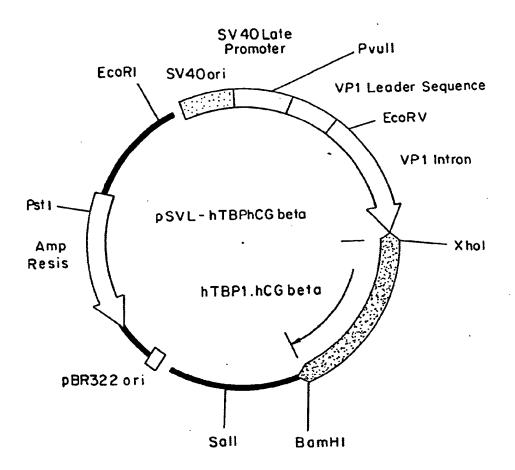
CCC ACT CCA CTA AGG TCC AAG AAG ACG ATG TTG GTC CAA AAG AAC GTC ACT TCA GAG TCC ACT TGC TGT GTA GCT AAA TCA TAT AAC AGG GTC Pro thr pro Leu Arg Ser Lys Lys thr Met Leu Val Gln Lys Asn Val thr Ser Glu Ser thr Cys Cys Val Ala Lys Ser Tyr Asn Arg Val

ACA GTA ATG GGG GGT TTC AAA GTG GAG AAC CAC ACG GCG TGC CAC TGC AGT ACT TGT TAT TAT CAC AAA TCT TAA Thr Val Met Gly Gly Phe Lys Val Glu Asn His Thr Ala Cys His Cys Ser Thr Cys Tyr Tyr His Lys Ser ...

Bam HI



FIG. 1b(1)





F1G.16(2)



nGH Signal Sequence

hGH Intron

CTCGAG ATG GCT ACA G GIAAGCGCCCTAAAAICCCTITGGGCACAAIGTGTCCTGAGGGAGGGTAGCGACTGTAGAIGGGACGGGGGGCACTAACCCICAGGIITGGG

PSer Arg Thr Ser Leu Leu Leu Ala Phe Gly Leu Leu Cys CTCTTGCTCCGGCTCCCTCTGTTGCCCTCTGGTTTCTCCCCAGGC

GAT AGT GTG TGT CCC CAA GGA AAA TAT ATC CAC CCT CAA AAT AAT TCG ATT TGC TGT ACC Asp Ser Val Cys Pro Gln Gly Lys Tyr Ile His Pro Gln Asn Asn Ser Ile Cys Cys Thr +20 Asp of Processed TBP1 CCC TGG CTT CAA GAG GGC AGT GCC Pro Trp Leu Gln Glu Gly Ser Ala

ANG TGC CAC AAA GGA ACC TAC TTG TAC AAT GAC TGT CCA GGC CCG GGG CAG GAT ACG GAC TGC AGG GAG TGT GAG AGC GGC TCC TTC ACC LYS CYS His Lys Gly Thr Tyr Leu Tyr Asn Asp Cys Pro Gly Pro Gly Gln Asp Thr Asp Cys Arg Glu Cys Glu Ser Gly Ser Phe Thr

GCT TCA GAA AAC CAC CTC AGA CAC TGC CTC AGC TGC TCC AAA TGC CGA AAG GAA ATG GGT CAG GTG GAG ATC TCT TGC ACA GTG GAC Ala Ser Glu Asn His Leu Arg His Cys Ieu Ser Cys Ser Lys Cys Arg Lys Glu Met Gly Gln Val Glu Ile Ser Ser Cys Thr Val Agp •

CGG GAC ACC GTG TGT GGC TGC AGG AAG AAC CAG TAC CGG CAT TAT TGG AGT GAA AAC CTT TTC CAG TGC TTC AAT TGC AGG CTC TGC CTC AAG AAG AAG LY8 AAA AAA GIN TYY AAG HI8 TYY TYP SET GIU AAN LEU PHE GIN CY8 PHE AAN CY8 SEY LEU CY8 LEU

AAT GGG ACC GTG CAC CTC TCC TGC CAG GAG AAA CAG AAC ACC GTG TGC ACC TGC CAT GCA GGT TTC TTT CTA AGA GAA AAT GAG TGT GTC ABN Gly Thr Val His Leu Ser Cys Glu Asn Glu Lys Gln Asn Thr Val Cys Thr Cys His Ala Gly Phe Phe Leu Arg Glu Asn Glu Cys Val +7 Pro of hCG beta Linker

TCC TGT GCT GGT GGT CCA CGG TGC CGC CCC ATC AAT GCC ACC CTG GCT GTG GAG AAG GAG GGC TGC CCC GTG TGC ATC ACC GTC Ser Cys Ala Gly Ala Gly Pro Arg Cys Arg Pro Ile Aen Ala Thr Leu Ala Val Glu Lys Glu Gly Cys Pro Val Cys Ile Thr Val AAC ACC AAC ATC TGT GCC GGC TAC TGC CCC ACC ATG ACC CGC GTG CTG CGG GTC CTG CCG GCC CTG CCT CAG GTG GTG TGC AAC TAC Asn Thr Thr ile Cys Ala Gly Tyr Cys Pro Thr Met Thr Arg Val Leu Gln Gly Val Leu Pro Ala Leu Pro Gln Val Val Cys Asn Tyr

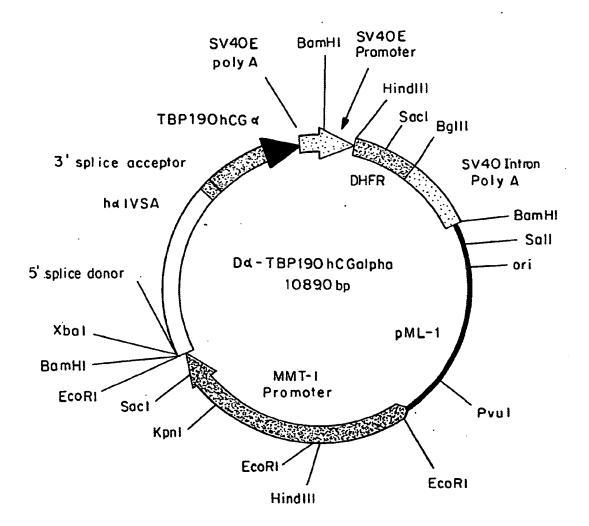
CGC GAT GTG CGC TTC GAG TCC ATC CGG CTC CCT GGC TGC CGG CGC GGC GTG AAC CCC GTG GTC TCC TAC GCC GTG GCT CTC AGC TGT CAA AKG AAB VA1 AKG Phe Glu Ser Ile Arg Leu Pro Gly Cys Pro Arg Gly Val Asn Pro Val Val Ser Tyr Ala Val Ala Leu Ser Cys Gln

TCC TCA AAG GCC CCT CCC CCC AGC CTT CCA AGC CCA TCC CGA CTC CCG GGG CCC TCG GAC ACC CCG ATC CTC CCA CAA TAA Ser Ser Lys Ala Pro Pro Pro Ser Leu Pro Ser Pro Ser Asp Thr Pro Ile Leu Pro Gln ***

Bam HI



FIG. 2a(1)





F16.20(2)

GTAAGCGCCCTTAAAATCCCTTTGGGCACAATGTGCTGAGGGGAGGAGGCAGCGACCTGTAGATGGGACGGGGGCACTAACCCTCAGGTTTGGGGTTTTCT hGH Intron hGH Signal Sequence TCGAG ATG GCT ACA G

CONCINCULARIZACION DE TOCIONE DE LOS LOS LOS CONTROLOS CON

CAA GAG GGC AGT GCC GAT AGT GTG TGT CCC CAA GGA AAA TAT ATC CAC CCT CAA AAT AAT TGG ATT TGC TGT ACC AAG TGC CAC AAA GGA GIN GIU GIY Ser Ala Asp Ser Val Cys Pro Gln Gly Bro Gln Asn Asn Ser IIe Cys Thr Lys Cys His Lys Gly +20 Asp of processed TBP1

ACC TAC TTG TAC AAT GAC TGT CCA GGC CCG GGG CAG GAT ACG GAC TGC AGG GAG TGT GAG AGC GGC TCC TTC ACC GCT TCA GAA AAC CAC CTC Thr Tyr Leu Tyr Asn Asp Cys Pro Gly Pro Gly Gln Asp Thr Asp Cys Arg Glu Cys Glu Ser Gly Ser Phe Thr Ala Ser Glu Asn His Leu

AGA CAC TGC CTC AGC TGC TCC AAA TGC CGA AAG GAA ATG GGT CAG GTG GAG ATC TCT TGC ACA GTG GAC CGG GAC ACC GTG TGT GGC TGC

AGG AAG AAC CAG TAC CGG CAT TAT 199 AGT GAA AAC CTT TTC CAG TGC TTC AAT 19C AGC CTC 19C CTC AAT GGG ACC GTG CAC TCC 19C AAG AAS GIN TYX AKG HIS TYX TXP Sex Glu Asn Leu Phe Gin Cys Phe Asn Cys Sex Leu Cys Leu Asn Gly Thx Val His Leu Sex Cys

CAG GAG AAA CAG AAC ACC GTG TGC ACC TGC CAT GCA GGT TTC TTT CTA AGA GAA AAC GAG TGT GTC TGT AGT AAC TGT AAG AAA AGC CTG
GIN GIN Lys Gin Asn Thi Val Cys Thi Cys His Ala Gly Phe Phe Leu Arg Glu Asn Glú Cys Val Ser Cys Ser Asn Cys Lys Lys Ser Leu

GCC CCA GGT TGC CCA +7 Cys of hCG alpha GAG TGC ACG AAG TTG TGC CTA CCC CAG ATT GAG AAT GTT AAG GGC ACT GAG GAC TCA GGC ACC ACA GCC GGT GCT Glu Cys Thr Lys Leu Cys Leu Pro Gln Ile Glu Aan Val Lys Gly Thr Glu Asp Ser Gly Thr Thr Ala Gly Ala Linker

GAA TGC ACG CTA CAG GAA AAC CCA TTC TTC TCC CAG CCG GGT GCC CCA ATA CTT CAG TGC ATG GGC TGC TTC TCT AGA GCA TAT CCC ACT Glu Cys Thx Leu Gln Glu Asn Pxo Phe Phe Ser Gln Pro Gly Ala Pro Ile Leu Gln Cys Met Gly Cys Cys Phe Ser Arg Ala Tyr Pro Thr

CCA CTA AGG TCC AAG AAG ACG ATG TTG GTC CAA AAG AAC GTC ACC TCA GAG TCC ACT TGC TGT GTA GCT AAA TCA TAT AAC AGG GTC ACA GTA

Pro Leu Arg Ser Lys Lys Thr Met Leu Val Gln Lys Asn Val Thr Ser Glu Ser Thr Cys Cys Val Ala Lys Ser Tyr Asn Arg Val Thr Val

ATG GGG GGT TIC AAA GIG GAG AAC CAC ACG GCG IGC CAC IGC AGT ACT IGT IAT IAT CAC AAA ICT IAA GGAICCCICGAG Met Gly Gly Phe Lys Val Glu Asn His Thr Ala Cys His Cys Ser Thr Cys Tyr Hys Lys Ser ***

Bam HI Xhol



FIG. 2b(1)

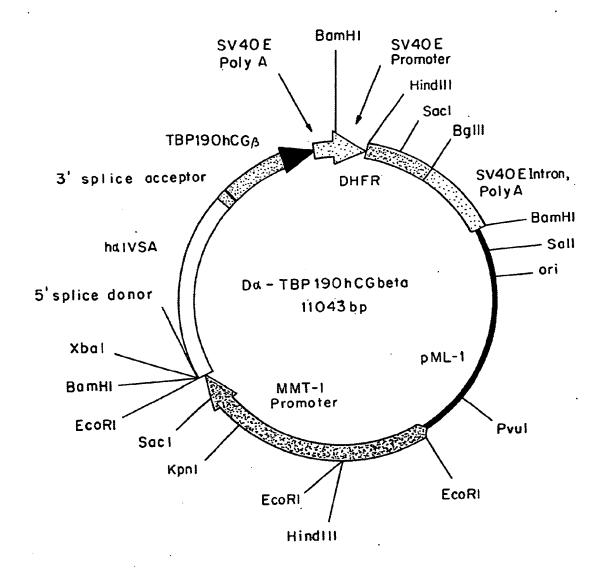


FIG. 2612,

ATG GCT ACA G GTAAGGGCCCCTTAAAATCCCTTTGGGCACAATGTGTCCTGAGGGGAGAAGCAGCGACGTAGATGGGAACGGGGGGCACTAACCCTCAGGTTTGGG hGH Intron hGH Signal Sequence ▶Met Ala Thr CTCGAG

r S 9 1 2 3 A) a Cig CTG CTC Leu Leu Ser CICTIBGACTOCOGGACCOCOTOTOGGATATCOCCCAGG C TCC CGG ACG

gat agt gtg tot ccc caa gga aaa tat atc cac cct caa aat aat tog att tgc tgt acc +20 Asp of Processed TBP1

CCC TGG CTT CAA GAG GGC AGT GCC PPro Trp Leu Gln Glu Gly Ser Ala

TTC ACC Amp Ser Val Cym Pro Gln Gly Lym Tyr Ile Him Pro Gln Amn Amn Ser Ile Cym Thr TCC ANG TGC CAC ANA GGA ACC TAC TTG TAC ANT GAC TGT CCA GGC CCG GGG CAG GAT ACG GAC TGC AGG GAG TGT GAG AGC GGC '

Lys Cys His Lys Gly Thr Tyr Leu Tyr Asn Asp Cys Pro Gly Pro Gly Gln Asp Thr Asp Cys Arg Glu Cys Glu Ser Gly

GAC ACA GTG Thr Val GAG ATC TCT TCT TGC Glu Ile Ser Ser Cys CAC CTC AGA CAC TGC CTC AGC TGC TCC ADA TGC CGA AAG GAA ATG GGT CAG GTG His Leu Arg His Cys Leu Ser Cys Ser Lys Cys Arg Lys Glu Met Gly Gln Val GLA AAC GCT TCA

760 ည TTC AAT TGC AGC Phe Asn Cys Ser COO GAC ACC OTO TOT GGC TGC AGG AAG AAC CAG TAC CGG CAT TAT TGG AGT GAA AAC CTT TTC CAG TGC 'AKG ABP The Val Cys Gly Cys Akg Lys Asn Gln Tyr Arg His Tyr Tep Ser Glu Asn Leu Phe Gln Cys AAT GGG ACC GTG CAC CTC TGC CAG GAG AAA CAG ACC ACC GTG TGC CAT GCA GGT TTC TTT CTA AGA GAA AAC GAG TGT GTC AAB GJY Thr Val His Leu Ser Cys Gln Glu Lys Gln Asn Thr Val Cys Thr Cys His Ala Gly Phe Phe Leu Arg Glu Asn Glu Cys Val

TCC TGT AGT AAC TGT AAG AAA AGC CTG GAG TGC ACG AAG TTG TGC CTA CCC CAG ATT GAG AAT GTT AAG GGC ACT GAG GAC TCA GGC ACC 'Ser Cys Ser Asn Cys Lys Lys Ser Glu Cys Thr Glu Asp Ser Gly Thr +7 Pro of beta

ACA GCT GGT GCT CCA CGG TGC CGC CCC ATC AAT GCC ATC TG GCT GTG GAG AAG GAG GGC TGC CCC GTG TGC ATC ACC GTC AAC

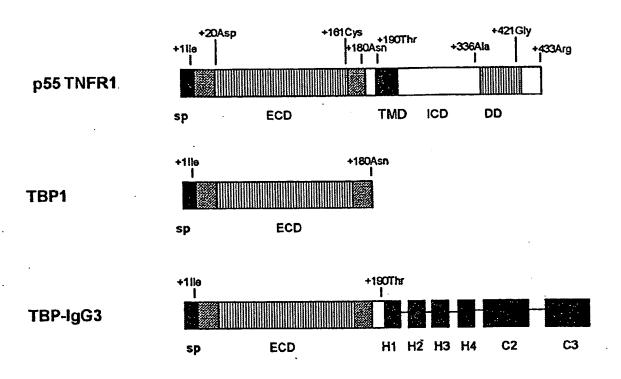
CAG GTG GTG TGC AAC TAC CGC Gln Val Val Cys Asn Tyr Arg ACC ACC ATC TGT GCC GGC TAC TGC CCC ACC ATG ACC CGC GTG CTG CAG GGG GTC CTG CCG GCC CTG CCT FThr Thr Ile Cyg Ala Gly Tyr Cyg Pro Thr Met Thr Arg Val Leu Gln Gly Val Leu Pro Ala Leu Pro GAG TCC ATC CGG CTC CCT GGC TGC CCG GGC GTG AAC CCC GTG GTC TCC TAC GCC GTG GCT CTC AGC TGT CAA TGT Glu Ser Ile Arg Leu Pro Gly Cys Pro Arg Gly Val Asn Pro Val Val Ser Tyr Ala Val Ala Leu Ser Cys Gln Cys GAT GTG CGC TTC Asp Val Arg Phe

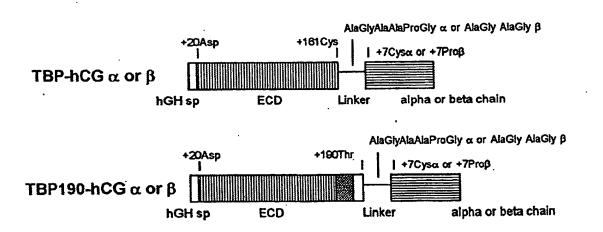
GCA CTC TGC CGC CGC AGC ACT GAC TGC GGG GGT CCC AAG GAC CAC CCC TTG ACC TGT GAT/GAC CCC CGC TTC CAG GAC TCC TCT TCC AAL Leu Cys Arg Arg Ser Thr Thr Asp Cys Gly Gly Pro Lys Asp His Pro Leu Thr Cys Asp Asp Pro Arg Phe Gln Asp Ser Ser Ser Ser

TER ANG GCC CET CCC CCC AGC CTT CCA AGC CCA TCC CGA CTC CCG GGG CCC TCG GAC ACC CCG ATC CTC CCA CAA TAA GGATCCCTCGAG P Ser Lys Ala Pro Pro Ser Leu Pro Ser Pro Ser Arg Leu Pro Gly Pro Ser Asp Thr Pro Ile Leu Pro Gln ***

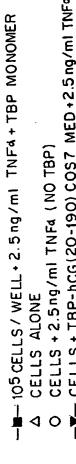


FIG. 3
p55 TNFR1, TBP1 and TBP1 FUSION CONSTRUCTS



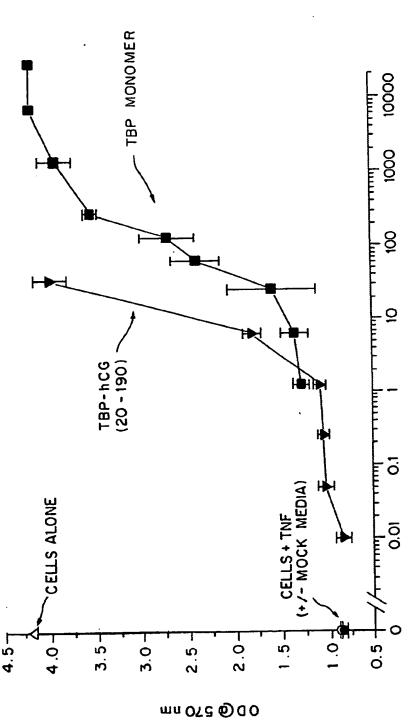


F16.4

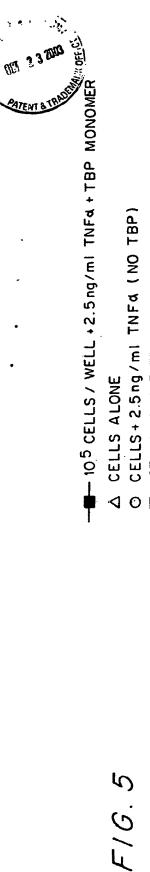


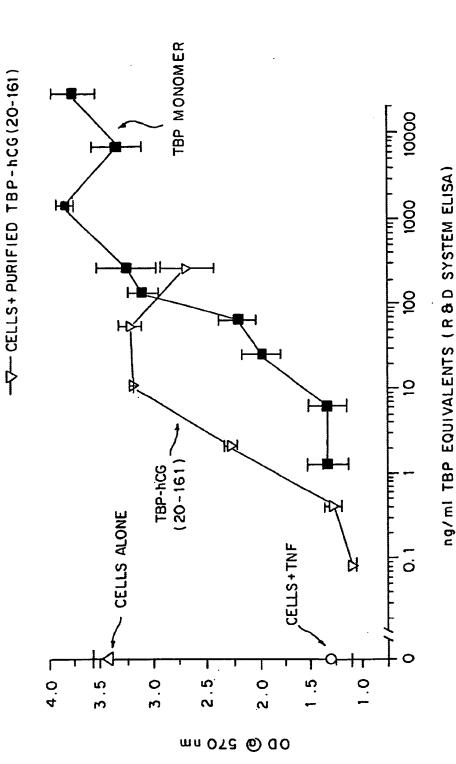
CELLS + TBP-hCG(20-190) COS7 MED+2.5 ng/ml TNF4

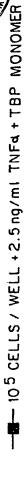
CELLS + COS7 MOCK TRANSFECTANT MEDIA + 2.5 ng/ml TNF4



ng /mi TBP EQUIVALENTS(R&D SYSTEM ELISA)





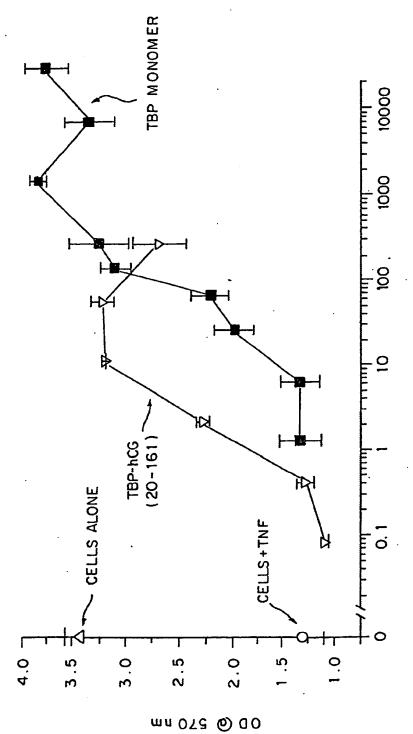


A CELLS ALONE

F16.6

O CELLS + 2.5 ng / ml TNF4 (NO TBP)

-Q- CELLS+ PURIFIED TBP-hCG (20-161)



ng/ml TBP EQUIVALENTS (RBD SYSTEM ELISA)